DOCKET NO.: ISIS-5325 PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In Re Application of:

Charles Allerson et al. Confirmation No.: 5641
Application No.: 10/701,007
Group Art Unit: 1635

For: Compositions Comprising Alternating 2'-Modified Nucleosides For Use In Gene

Modulation

Filing Date: November 4, 2003

ELECTRONICALLY FILED: September 16, 2009

Examiner: Jane J. Zara

Mail Stop Amendment Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Pursuant to 37 CFR § 1.56 and in accordance with 37 CFR §§ 1.97-1.98, information relating to the above-identified application is hereby disclosed. Inclusion of information in this statement is not to be construed as an admission that this information is material as that term is defined in 37 CFR § 1.56(b).

In accordance with § 1.97(b), since this Information Disclosure Statement is being filed either within three months of the filing date of the above-identified application, within three months of the date of entry into the national stage of the above identified application as set forth in § 1.491, before the mailing date of a first Office Action on the merits of the above-identified application, or before the mailing date of a first Office Action after the filing of request for continued examination under § 1.114, no additional fee is required.

\boxtimes	In accordance with § 1.97(c), this Information Disclosure Statement is being
	filed after the period set forth in § 1.97(b) above but before the mailing date of
	either a Final Action under § 1.116 or a Notice of Allowance under § 1.311, or
	before an action that otherwise closes prosecution in the application, therefore:
	Certification in Accordance with § 1.97(e) is attached; or
	The fee of $$180.00$ as set forth in $$1.17(p)$ is attached.
	In accordance with § 1.97(d), this Information Disclosure Statement is being
	filed after the mailing date of either a Final Action under § 1.113 or a Notice
	of Allowance under § 1.311 but before, or simultaneously with, the payment
	of the Issue Fee, therefore included are: Certification in Accordance with §
	1.97(e); and the submission fee of §180.00 as set forth in § 1.17(p).
\boxtimes	Copies of reference numbers 362-979 listed on the attached Form PTO-1449

are enclosed herewith.

Copies of reference numbers **1-361** on the attached Form PTO 1449 are not required to be submitted pursuant to 37 CFR § 1.98(a)(2)(ii).

The enclosed 1449 form includes references cited in related applications. For the Examiner's convenience, also being submitted herewith is a table listing related applications and references cited in official actions issued in the related applications that served as the basis for rejections under 35 U.S.C. § 102 or § 103.

The relevance of those listed references which are not in the English language is as follows:

English language abstracts have been provided for those listed references which are not in the English language.

Please charge any deficiency or credit any overpayment to Deposit Account No. 23-3050.

Date: September 16, 2009 /Jane E. Inglese/ Jane E. Inglese

Registration No. 48,444

WOODCOCK WASHBURN LLP Cira Centre 2929 Arch Street, 12th Floor Philadelphia, PA 19104-2891 Telephone: (215) 568-3100

Facsimile: (215) 568-3439

REJECTIONS MADE IN OTHER APPLICATIONS

Application Number/ Attorney	Date of Official Action on the	Rejections Levied in Official Action under 35 U.S.C. §§	Other Rejections Levied in Official Action	Application Status
Docket Number	Merits	102 or 103	III OIIICIAI ACCIOII	Status
08/659,440 ISIS2197	January 13, 1997	§103 (a): Metelev, Bioorg. Med. Chem. Lett. 1994, 4:2929-2934; and Lengyel Journal of Interferon Res., 1987, 7, 511	§112, first paragraph, enablement	Patented
	July 22, 1997	§ 103 (a): Strickland, <i>Science</i> 1988 241:680-684; Metelev, <i>Bioorg. Med. Chem.</i> <i>Lett.</i> 1994, 4:2929-2934; and Dagle, <i>Nucleic Acids Res.</i> 1991, 19, 1805-1810		
	January 28, 1998	§ 103 (a): Strickland, <i>Science</i> 1988 241:680-684; Metelev, <i>Bioorg. Med. Chem.</i> <i>Lett.</i> 1994, 4:2929-2934; and Goodchild, Bioconjug. Chem. 1990 1:165-187		
08/870,608 ISIS2484	March 17, 1999	None	(1) Obviousness-type double patenting U.S. patent application no. 08/659,440; (2) § 112 first paragraph, enablement	Patented

	October 8, 1999	None	(1) Obviousness-type double patenting U.S. patent application no. 08/659,440; (2) § 112 first paragraph, enablement	
09/479,783 ISIS4313	May 1, 2003	§ 102 (b) PCT patent application publication no. WO 94/01550		Abandoned
	December 24, 2003	§ 102 (b) PCT patent application publication no. WO 94/01550		
	May 18, 2004	(1) § 102 (b) U.S. patent no. 5,013,830; (2) § 102 (b) U.S. patent no. 5,256,775	(1) § 112, first paragraph, written description; (2) § 112, second paragraph indefiniteness	
	February 9, 2005	(1) § 102 (b) U.S. patent no. 5,013,830; (2) § 102 (b) U.S. patent no. 5,256,775	§ 112, first paragraph, written description	
	September 1, 2005	(1) § 102 (b) U.S. patent no. 5,013,830; (2) § 102 (b) or § 103(a) U.S. patent no. 5,256,775	(1) § 112, first paragraph, written description; (2) § 112, second paragraph indefiniteness	
	June 8, 2006	(1) § 102 (b) U.S. patent no. 6,849,726; (2) § 103 (a) U.S. patent no. 6,849,726		
	February 21, 2007	§ 101, utility		

10/280,600 ISIS0002-104	March 28, 2006		(1) § 112, first paragraph, written description; (2) § 112, second paragraph indefiniteness	Abandoned
10/281,349 ISIS0002-105 (ISIS-5780)	June 30, 2006	(1) § 103 (a) U.S. patent no. 6,573,072; (2) §103(a) U.S. patent no. 6,849,726	§ 112, second paragraph indefiniteness	Allowed
	March 19, 2007	§ 103 (a) U.S. patent no. 6,573,072	Obviousness-type double patenting, U.S. patent no. 6,107,094	
	December 11, 2007	§ 103 (a) U.S. patent no. 6,087,484	(1) § 112, second paragraph indefiniteness; (2) § 112, first paragraph written description	
	June 19, 2009		Obviousness-type double patenting, U.S. patent no. 7,432,250	
10/281,312 ISIS0002-106 (ISIS-5779)	June 29, 2006	§ 102 (e) U.S. patent no. 6,573,072	§ 112, second paragraph indefiniteness	Patented
	March 10, 2008		Obviousness-type double patenting, U.S. patent no. 6,107,094	
10/281,297 ISIS0002-107 (ISIS-5778)	March 21, 2006	§ 102 (b) PCT patent application publication no. WO 94/01550		Patented
	September 22, 2006	(1) § 103 (a) U.S. patent no. 6,573,072; (2) §103(a) U.S. patent no.		

		6,849,726		
	April 2, 2007	§ 103 (a) U.S. patent no. 6,573,072	Obviousness-type double patenting, U.S. patent no. 6,107,094	
10/078,949 ISIS5027	February 10, 2005	(1) § 102 (b) PCT patent application publication no. WO 94/01550; (2) § 103 (a) PCT patent application publication no. WO 94/01550 in view of Hunzinker and Leumann, <i>Nucleic Acid Analogues: Synthesis and Properties in Modern Synthetic Methods</i> , 1995, ed. Ernst and Leumann, pp. 331-417	§ 112, first paragraph, written description	Pending
	August 12, 2005		(1) § 112, second paragraph indefiniteness; (2) § 112, first paragraph, written description	
	June 23, 2006	(1) § 102 (e) U.S. patent no. 6,573,072; (2) §102 (e) U.S. patent no. 6,849,726; (3) § 103 (a) U.S. patent no. 6,573,072; (4) §103(a) U.S. patent no. 6,849,726; (5) § 103 (a) U.S. patent no. 6,573,072; (6) § 103 (a) U.S. patent no.		

	May 23, 2008	6,573,072 in view of U.S. patent no. 6,037,463; (7) §103(a) U.S. patent no. 6,849,726 in view of U.S. patent no. 6,037,463 §103(a) U.S. patent no. 6,087,484	(1) § 112, second paragraph indefiniteness; (2) § 112, first paragraph, written description	
	September 11, 2008	§ 103 (a) PCT patent application publication no. WO 94/02498	Obviousness-type double patenting, (1) U.S. patent no. 6,107,094; (2) U.S. patent no. 5,898,031; (3) U.S. patent application no. 10/281,349	
10/371,526 ISIS0002-108	December 13, 2005	(1) § 102 (b) European patent no. EP 0 339 842; (2) § 103 (a) European patent no. EP 0 339 842 in view of Milligan, <i>J. Med. Chem</i> , 1993, 36, 1923; PCT patent application publication no. WO 93/07883; and U.S. patent no. 5,898,031	(1) § 112, second paragraph indefiniteness; (2) § 112, first paragraph, written description; (3) § 112, first paragraph, enablement	Abandoned
10/860,455 CHEM0003US.P2 (ISIS-5480)	March 23, 2007	(1) § 102 (a) Tracewell, Toxicology and Applied Pharmacology, 1995, 135, 179- 184; (2) § 102 (b) PCT patent application publication no. WO	Obviousness-type double patenting, U.S. patent application no. 09/479,783	Pending

		94/01550		
	October 31, 2007	(1) § 102 (a) Tracewell, Toxicology and Applied Pharmacology, 1995, 135, 179- 184; (2) § 103(a) US 5,506,212	Obviousness-type double patenting, U.S. patent application no. 09/479,783	
	July 18, 2008	(1) § 103(a) U.S. patent application publication no. 2004/0259247 in view of U.S. patent no. 5,506,212; (2)	Obviousness-type double patenting, U.S. patent application no. 09/479,783	
10/701,012 CHEM0004US.P1	May 8, 2006	(1) § 102 (e) U.S. patent application publication no. 2003/0139585; (2) § 102 (e) U.S. patent application publication no. 2004/0146867; (3) § 103 (a) U.S. patent application publication nos. 2003/0139585 and 2004/0146867 in view of U.S. patent nos. 5,082,934 and 5,719,271	(1) § 112, second paragraph indefiniteness; (2) § 112, first paragraph, written description; (3) § 112, first paragraph, enablement; (4) Obviousness-type double patenting, U.S. patent application no. 10/606,510	Abandoned

10/700,884	June 24, 2005	(1) § 102 (b) Parrish,	Obviousness-type double	Abandoned
CHEM0008US.P1		Molecular Cell, 2000, 6, 1077-	patenting, U.S. patent	
(ISIS-5317)		1087;	application no. 10/700,884	
		(2) § 102 (b) U.S. patent no.		
		5,891,684		
	July 12, 2006	(1) § 102 (e) U.S. patent no.	(1) 101, non-statutory	
		6,222,025;	subject matter;	

			Nucleic Acids Res. 1994, 22, 1429-1436, and Martin, Helvetica Chimica Acta, 1995, 78, 486-504;	
			· · · · · · · · · · · · · · · · · · ·	
			patent no. 5,760,202 in view of U.S. patent no.	
			5,256,775, U.S. patent no. 5,466,786, Kuimelis, <i>Nucleic Acids Res.</i> 1994,	
			22, 1429-1436, and Martin, Helvetica Chimica Acta,	
10/700,939	May 25, 2006	(1) § 102 (b) U.S. patent no.	1995, 78, 486-504; (1) Claim of priority	Abandoned

CHEM0012US.P1 (ISIS-5318)		5,561,043; (2) § 102 (b) U.S. patent no. 5,424,413; (3) § 102 (b) U.S. patent no. 6,274,723; (4) § 102 (b) Porta & Lizardi, <i>Biotechnology</i> , 1994, 13, 161- 164	denied; (2) § 112, first paragraph, enablement	
	July 28, 2006	(1) § 102 (b) U.S. patent no. 5,561,043; (2) § 102 (b) U.S. patent no. 5,424,413; (3) § 102 (b) U.S. patent no. 6,274,723; (4) § 102 (b) Porta & Lizardi, <i>Biotechnology</i> , 1994, 13, 161-164	(1) Claim of priority denied; (2) § 112, first paragraph, enablement	
10/701,316 ISIS5301	October 13, 2006	(1) § 102 (b) U.S. patent no. 5,998,203; (2) § 102 (b) PCT patent application publication no. WO 94/01550; (3) § 102 (b) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888	(1) Claim of priority denied; (2) § 112, first paragraph, enablement	Pending
	March 6, 2007		§ 112, first paragraph, written description	
	July 10, 2007	(1) § 103 (a) Elbashir, <i>EMBO</i> J., 2001, 20, 6877-6888 in view of Wilds, <i>Nucleic Acids Res.</i> , 2000, 28, 3625-3635 and Hammond, <i>Nature</i> , 2001, 2,	(1) Claim of priority denied; (2) § 112, first paragraph, written description	

		110-119; (2) § 103 (a) Elbashir, EMBO J., 2001, 20, 6877-6888 in view of Wilds, Nucleic Acids Res., 2000, 28, 3625-3635, Hammond, Nature, 2001, 2, 110-119, and Veronese, Il Farmaco, 1999, 54, 497-516; (3) § 103 (a) Tracewell, Toxicology and Applied Pharmacology, 1995, 135, 179- 184 in view of Wilds, Nucleic Acids Res., 2000, 28, 3625- 3635		
	March 27, 2008	§ 102 (e) U.S. patent application publication no. 2006/0127891	(1) Claim of priority denied; (2) § 112, second paragraph, indefiniteness; (3) § 112, first paragraph, written description	
	October 30, 2008	§ 103 (a) U.S. patent application publication no. 2006/0127891 in view of U.S. patent application publication no. 2004/0180351 and Olie, et al., Biochimica et Biophysica Acta, 2002, 1576, 101-109	(1) Claim of priority denied; (2) § 112, second paragraph, indefiniteness; (3) § 112, first paragraph, written description	
10/700,689 ISIS5313	May 31, 2005	(1) § 102 (b) Parrish, Molecular Cell, 2000, 6, 1077- 1087;	Obviousness-type double patenting, U.S. patent application no. 10/701,316	Pending

		(2) § 102 (a) U.S. patent no.		
-	November 29, 2005	5,998,203 (1) § 102 (b) Parrish,		
	11070111001 25, 2000	Molecular Cell, 2000, 6, 1077-		
		1087;		
		(2) § 103 (a) Parrish,		
		Molecular Cell, 2000, 6, 1077-		
		1087 in view of Elbashir,		
		EMBO J., 2001, 20 (23), 6877-		
		6888, U.S. patent no. 5,955,443		
		and Hammond, <i>Nature</i> , 2001, 2, 110-119		
	July 28, 2006	(1) § 102 (b) Parrish,	(1) Claim of priority	
	July 20, 2000	Molecular Cell, 2000, 6, 1077-	denied;	
		1087;	(2) Obviousness-type	
		(2) § 103 (a) Parrish,	double patenting, U.S.	
		Molecular Cell, 2000, 6, 1077-	patent application no.	
		1087 in view of Elbashir,	10/701,316;	
		EMBO J., 2001, 20 (23), 6877-	(3) § 112, second	
		6888, U.S. patent no. 5,955,443	paragraph indefiniteness	
		and Hammond, <i>Nature</i> , 2001,		
_	1 14 2006	2, 110-119	(1) (1) (1)	
	November 14, 2006	(1) § 102 (b) Parrish,	(1) Claim of priority	
		Molecular Cell, 2000, 6, 1077-	denied;	
		1087; (2) § 103 (a) Parrish,	(2) Obviousness-type double patenting, U.S.	
		Molecular Cell, 2000, 6, 1077-	patent application no.	
		1087 in view of Elbashir,	10/701,316	
		EMBO J., 2001, 20 (23), 6877-		
		6888, U.S. patent no. 5,955,443		
		and Hammond, <i>Nature</i> , 2001,		

	A 110 110	
	2,110-119;	
	(3) § 102 (b) U.S. patent no.	
	5,998,203	
September 18, 2007	(1) § 102 (e) U.S. patent no.	(1) § 112, first paragraph,
	7,022,828;	written description;
	(2) § 103 (a) Elbashir, <i>EMBO</i>	(2) Obviousness-type
	J., 2001, 20, 6877-6888 in	double patenting, U.S.
	view of Wilds, Nucleic Acids	patent application no.
	Res., 2000, 28, 3625-3635,	10/701,316;
	Parrish, Molecular Cell, 2000,	(3) Obviousness-type
	6, 1077-1087, Monia, <i>J Biol</i> .	double patenting, U.S.
	Chem., 1993, 268, 14514-	patent no. 6,107,094;
	14522 and Hammond, Nature	(4) Obviousness-type
	Reviews Genetics, 2001, 2,	double patenting, U.S.
	110-119;	patent application no.
	(3) § 103 (a) Bevilacqua,	10/281,297;
	Biochemistry, 1996, 35, 9983-	(5) Obviousness-type
	9994 in view of Monia, J Biol.	double patenting, U.S.
	Chem., 1993, 268, 14514-	patent application no.
	14522 and U.S. patent no.	10/078,949;
	5,631,148	(6) Obviousness-type
		double patenting, U.S.
		patent application no.
		10/860,265;
		(7) Obviousness-type
		double patenting, U.S.
		patent application no.
		10/701,007
June 17, 2008	§ 102 (e) U.S. patent	(1) Claim of priority
	application publication no.	denied;
	2004/0180351	(2) Obviousness-type

			double patenting, U.S. patent application no. 10/701,316; (3) Obviousness-type double patenting, U.S. patent no. 6,107,094; (4) Obviousness-type double patenting, U.S. patent application no. 10/860,265; (5) Obviousness-type double patenting, U.S. patent application no. 10/701,007; (6) § 112, first paragraph, written description;	
	January 16, 2009	§ 103 (1) U.S. patent application publication no. 2004/0180351 in view of U.S. patent application publication no. 2006/0127891 and Olie, et al., Biochimica et Biophysica Acta, 2002, 1576, 101-109	(1) Claim of priority denied; (2) § 112, first paragraph, written description;	
10/701,264 ISIS5314	October 5, 2006	(1) § 102 (a) Elbashir, <i>EMBO</i> J., 2001, 20, 6877-6888; (2) § 102 (b) PCT patent application publication no. WO 94/01550; (3) § 102 (b) Monia, <i>J Biol.</i> Chem., 1993, 268, 14514-	(1) Claim of priority denied; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,316; (3) Obviousness-type	Pending

	14522; (4) § 102 (b) Yu, <i>Bioorganic</i> and Medicinal Chemistry, 1996, 4, 1685-1692; (5) § 102 (b) PCT patent application publication no. WO 94/02498	double patenting, U.S. patent application no. 10/701,265	
March 7, 2007	(1) § 102 (b) Shuman, <i>J. Biol Chem</i> , 1993, 268, 18943-18950; (2) § 103 (a) Beigelman, <i>J. Biol Chem</i> , 1995, 270, 25702-25708 in view of Koizumi, <i>Nucleic Acids Research</i> , 1989, 17, 7059-7071	(1) Obviousness-type double patenting, U.S. patent application no. 10/701,265; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,316; (3) Obviousness-type double patenting, U.S. patent application no. 09/479,783; (4) § 112, first paragraph, enablement	
July 25, 2007	(1) § 102 (a) Bevilacqua, Biochemistry, 1996, 35, 9983- 9994; (2) § 102 (a) Yu, RNA, 1997, 324-331; (3) § 103 (a) Beigelman, J. Biol Chem, 1995, 270, 25702-25708 in view of Koizumi, Nucleic Acids Research, 1989, 17, 7059-7071 and U.S. patent no.	(1) Obviousness-type double patenting, U.S. patent application no. 10/701,265; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,316; (3) Obviousness-type double patenting, U.S.	

February 6, 2008	5,151,510; (4) § 103 (a) Yu, <i>RNA</i> , 1997, 324-331 in view of U.S. patent no. 5,151,510 § 103 (a) U.S. patent application publication no. 2003/0143732 in view of Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888; Parrish, <i>Molecular Cell</i> , 2000, 6, 1077-1087; and U.S. patent no. 5,801,154.	patent application no. 09/479,783; (4) § 112, second paragraph, indefiniteness (1) Obviousness-type double patenting, U.S. patent application no. 10/701,265; (2) Obviousness-type double patenting, U.S. patent application no. 09/479,783; (3) Claim for priority denied	
August 29, 2008	§ 103 (a) U.S. patent application publication no. 2003/0143732 in view of Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888; Parrish, <i>Molecular Cell</i> , 2000, 6, 1077-1087; and U.S. patent no. 5,801,154.	(1) Obviousness-type double patenting, U.S. patent application no. 10/701,265; (2) Obviousness-type double patenting, U.S. patent application no. 09/479,783; (3) Claim for priority denied	
February 19, 2009	§ 103 (a) Lee, et al., Cell, 1993, 75, 843-854; Manche, et al., Molecular and Cellular Biology, 1992, 12, 5238-5248, published PCT application publication no. WO 94/01550, and U.S. patent no. 5,801,154		

11/054,848	March 23, 2007	(1) § 103 (a) Elbashir, <i>EMBO</i>	(1) Obviousness-type	Pending
ISIS5586	Water 25, 2007	J., 2001, 20, 6877-6888, U.S.	double patenting, U.S.	1 chang
15155500		patent application publication	patent application no.	
		no. 2003/014732, and U.S.	10/701,007;	
		patent application no.	(2) Obviousness-type	
		2003/0206887 in view of U.S.	double patenting, U.S.	
		patent no. 6,262,036, U.S.	patent application no.	
		patent application publication	10/860,265;	
		no. 2005/0142535, and U.S.	(3) § 112, first paragraph,	
		patent no. 6,133,246	enablement	
	November 30, 2007	§ 103 (a) Elbashir, <i>EMBO J</i> .,	(1) Obviousness-type	
	1101011001 30, 2007	2001, 20, 6877-6888, U.S.	double patenting, U.S.	
		patent application publication	patent application no.	
		no. 2003/014732, and U.S.	10/701,007;	
		patent application publication	(2) Obviousness-type	
		no. 2003/0206887 in view of	double patenting, U.S.	
		U.S. patent no. 6,262,036, U.S.	patent application no.	
		patent application publication	10/860,265	
		no. 2005/0142535, and U.S.	,	
		patent no. 6,133,246		
	June 18, 2008	§ 103 (a) Elbashir, EMBO J.,	(1) Obviousness-type	
		2001, 20, 6877-6888, U.S.	double patenting, U.S.	
		patent application publication	patent application no.	
		no. 2003/014732, and U.S.	10/701,007;	
		patent application no.	(2) Obviousness-type	
		2003/0206887 in view of U.S.	double patenting, U.S.	
		patent no. 6,262,036, U.S.	patent application no.	
		patent application publication	10/860,265	
		no. 2005/0142535, and U.S.		
		patent no. 6,133,246		
10/700,697	September 21, 2006	(1) § 103 (a) U.S. patent no.	(1) Claim of priority	Pending

ISIS5312		6,818,759 in view of U.S. patent no. 6,506,559; (2) § 103 (a) U.S. patent no. 6,506,559 in view of Alahari, <i>J. Pharmacology and Experimental Therapeutics</i> , 1998, 286, 419-428 and U.S. patent application publication	denied; (2) § 112, second paragraph, indefiniteness; (3) § 112, first paragraph, enablement; (4) § 112, first paragraph, written description; (5) Obviousness-type	
		no. 2003/0125241	double patenting, U.S. patent application no. 10/701,007	
	January 5, 2007	(1) § 103 (a) U.S. patent no. 6,818,759 and U.S. patent no. 6,506,559; (2) § 103 (a) U.S. patent no. 6,506,559, Alahari, <i>J. Pharmacology and Experimental Therapeutics</i> , 1998, 286, 419-428 and U.S. patent application publication no. 2003/0125241; (3) § 102 (e) Tuschl, <i>Molecular Interventions</i> , 2002, 2, 158-167	(1) Claim of priority denied; (2) § 112, first paragraph, written description; (3) Obviousness-type double patenting, U.S. patent application no. 10/701,007	
	July 6, 2007	(1) § 102 (b) Cook, Anti- Cancer Drug Design, 1991, 6, 585-607; (2) § 103 (a) Crooke, Biochemical Journal, 1995, 312, 599-608 in view of Berger, Nucleic Acids	(1) Claim of priority denied; (2) § 112, second paragraph, indefiniteness	

March 3, 2008	Research, 1998, 26, 2473-2480; (3) § 103 (a) Lesnik, Biochemistry, 1995, 34,10807-10815 in view of Berger, Nucleic Acids Research, 1998, 26, 2473-2480; (4) § 103 (a) Wu, J. Biol. Chem, 1998, 273, 2352-2542 in view of Cook, Anti-Cancer Drug Design, 1991, 6, 585-607 (1) § 102 (e) U.S. patent application publication no. 2004/019626; (2) § 102 (a) and § 102 (e) U.S. patent application publication no. 2003/0143732	(1) Claim of priority denied; (2) § 112, first paragraph, written description	
October 22, 2008	(1) § 103 (a) Grünweller, et al., Nucleic Acids Res., 2003, 31, 3185-3193 in view of U.S. patent application publication no. 2003/0143732; (2) § 103 (a) Braasch, et al., Biochemistry, 2003, 42, 7967-7975 in view of U.S. patent application publication no. 2003/0143732; (3) § 103 (a) published PCT application no. WO 2004/083430	(1) Claim of priority denied; (2) § 112, first paragraph, written description (3) Obviousness-type double patenting, U.S. patent application no. 10/701,264; (4) Obviousness-type double patenting, U.S. patent application no. 10/561,324	

	July 16, 2009	(1) § 103 (a) Grünweller, et al., Nucleic Acids Res., 2003, 31, 3185-3193 in view of U.S. patent application publication no. 2003/0143732; (2) § 103 (a) Braasch, et al., Biochemistry, 2003, 42, 7967-7975 in view of U.S. patent application publication no. 2003/0143732; (3) § 103 (a) published PCT application publication no. WO 2004/083430 (4) § 102 (e) U.S. patent application publication no. 2004/0180351	(1) § 112, first paragraph, written description; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,264; (3) Obviousness-type double patenting, U.S. patent application no. 10/561,324	
10/701,236 ISIS5207	October 2, 2006	(1) § 103 (a) PCT patent application no. WO 94/01550 in view of U.S. patent no. 5,639,873; (2) § 103 (a) U.S. patent application publication no. 2004/0029275 in view of U.S. patent no. 5,639,873	(1) Claim of priority denied; (2) § 112, first paragraph, written description; (3) § 112, first paragraph, enablement; (4) Obviousness-type double patenting, U.S. patent application no. 10/700,697	Pending
	March 14, 2007	(1) § 103 (a) U.S. patent application no. 2004/0029275 in view of U.S. patent no. 5,639,873		

(2) § 102 (a) Yu, RNA, 1997, 3, 324-331; (3) Obviousness-type double patenting, U.S. patent application publication no. 2004/0029275; (4) § 103 (a) Beigelman, J. Biol Chem, 1995, 270, 25702-25708 in view of Koizumi, Nucleic Acids Research, 1989, 17, paragraph, indefiniteness; (3) Obviousness-type double patenting, U.S. patent application no. 10/700,697; (4) Obviousness-type double patenting, U.S. patent application no. 10/701,264;	April 15, 2008	5,151,510; (5) § 103 (a) Yu, RNA, 1997, 3, 324-331 in view of U.S. patent no. 5,151,510 (1) § 102 (a) Bevilacqua, Biochemistry, 1996, 35, 9983-9994; (2) § 102 (a) Yu, RNA, 1997, 3, 324-331; (3) § 103 (a) Yu, RNA, 1997, 3, 324-331 in view of U.S. patent no. 5,151,510 and U.S. patent no. 5,142,047	double patenting, U.S. patent application no. 10/701,316 (1) Obviousness-type double patenting, U.S. patent application no. 10/700,697; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,264; (3) Obviousness-type double patenting, U.S. patent application no. 10/701,264;	
		324-331; (3) § 102 (e) U.S. patent application publication no. 2004/0029275; (4) § 103 (a) Beigelman, <i>J. Biol</i> <i>Chem</i> , 1995, 270, 25702-25708 in view of Koizumi, <i>Nucleic</i> <i>Acids Research</i> , 1989, 17, 7059-7071 and U.S. patent no.	(3) Obviousness-type double patenting, U.S. patent application no. 10/700,697; (4) Obviousness-type double patenting, U.S. patent application no. 10/701,264; (5) Obviousness-type	

		324-331 in view of U.S. patent no. 5,151,510 and U.S. patent no. 5,142,047	double patenting, U.S. patent application no. 10/700,697; (2) Obviousness-type double patenting, U.S. patent application no. 10/701,264; (3) Obviousness-type double patenting, U.S. patent application no. 10/701,316	
	April 2, 2009	§ 103 (a) Lee, et al., <i>Cell</i> , 1993, 75, 843-854; Manche, et al., <i>Mol. Cell Biol.</i> , 1992, 12, 5238-5248; PCT patent application publication no. WO 94/01550; U.S. patent no. 5,801,154; and U.S. patent no. 5,519,134		
10/700,920 ISIS5203	October 2, 2006	(1) § 102 (e) U.S. patent application publication no. 2004/0029275; (2) § 103 (a) U.S. patent application publication no. 2004/0029275 in view of U.S. patent no. 5,459,255; (3) § 103 (a) PCT patent application publication no. WO 94/01550 in view of U.S. patent no. 5,459,255	(1) Claim of priority denied; (2) § 112, first paragraph, written description; (3) § 112, first paragraph, enablement; (4) Obviousness-type double patenting, U.S. patent application no. 10/561,618	Abandoned

patent application no. 10/701,217; (7) Obviousness-type double patenting, U.S. patent application no. 10/701,236; (8) Obviousness-type double patenting, U.S. patent application no. 10/701,265 10/606,510 April 3, 2006 (1) §102 (e) U.S. patent (1) § 112, second Abandoned	10/460,433 CHEM0003US (ISIS-5200)	ruary 15, 2006	(1) § 102 (b) PCT patent application publication no. WO 94/01550; (2) § 102 (e) U.S. patent application publication no. 2003/0143732; (3) § 102 (b) U.S. patent no. 6,210,892; (4) § 102 (b) PCT patent application publication no. WO 02/44321; (5) U.S. patent application publication no. 2003/0143732 in view of U.S. patent no. 6,210,892	10/701,217; (7) Obviousness-type double patenting, U.S. patent application no. 10/701,236; (8) Obviousness-type double patenting, U.S.	Abandoned
--	---	----------------	--	--	-----------

CHEM0004US		application publication no. 2003/0139585; (2) §102 (e) U.S. patent application publication no. 2004/0146867; (3) §103 (a) U.S. patent application publication no. 2003/0139585 and U.S. patent application publication no. 2004/0146867 in view of U.S. patent no. 5,082,934 and U.S. patent no. 5,719,271	paragraph, indefiniteness; (2) § 112, first paragraph, written description; (3) § 112, first paragraph, enablement;	
10/701,285 CHEM0006US (ISIS-5240)	March 16, 2007		(1) § 112, first paragraph, written description; (2) § 101, utility and § 112, first paragraph, enablement	Pending
	August 30, 2007		(1) § 112, first paragraph, written description; (2) § 101, utility and § 112, first paragraph, enablement	
	April 11, 2008		(1) § 112, second paragraph indefiniteness (2) § 101, utility and § 112, first paragraph, enablement	
	December 24, 2008		§ 101, utility and § 112, first paragraph, enablement	
10/701,007 ISIS5325	May 5, 2006	§ 103 (a) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888, U.S. patent application publication no. 2003/0143732, and U.S. patent application publication		Pending

	no. 2003/0206887		
October 19, 2006	§ 103 (a) Elbashir, EMBO J.,		
	2001, 20, 6877-6888, U.S.		
	patent application publication		
	no. 2003/0143732, and U.S.		
	patent application publication		
	no. 2003/0206887		
March 26, 2007	§ 103 (a) Elbashir, <i>EMBO J.</i> ,	(1) § 112, first paragraph,	
	2001, 20, 6877-6888, U.S.	enablement;	
	patent application publication	(2) Obviousness-type	
	no. 2003/0143732, U.S. patent	double patenting, U.S.	
	application publication no.	patent application no.	
	2003/0206887, U.S. patent no.	10/860,265;	
	6,262,036, U.S. patent	(3) Obviousness-type	
	application publication no.	double patenting, U.S.	
	2005/0142535, and U.S. patent	patent application no.	
	no. 6,133,246	11/054,848	
C1 14 2007	0 102 (a) Ellas 1.1 EMDO I	(1) (1)	
September 14, 2007	§ 103 (a) Elbashir, <i>EMBO J.</i> ,	(1) Obviousness-type	
	2001, 20, 6877-6888, U.S.	double patenting, U.S.	
	patent application publication	patent application no.	
	no. 2003/0143732, U.S. patent	10/860,265;	
	application publication no.	(2) Obviousness-type	
	2003/0206887, U.S. patent no. 6,262,036, U.S. patent	double patenting, U.S.	
	' ' ' 1	patent application no.	
	application publication no.	11/054,848	
	2005/0142535, and U.S. patent		
March 27, 2000	no. 6,133,246	(1) Obviousness tyres	
March 27, 2008	§ 103 (a) Elbashir, <i>EMBO J.</i> ,	(1) Obviousness-type	
	2001, 20, 6877-6888, U.S.	double patenting, U.S.	
	patent application publication	patent application no.	

10/860,265 ISIS5482	April 10, 2007	§ 103 (a) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888, U.S.	(1) § 112, first paragraph, enablement;	Pending
10/0/0 0/5	4 110 2007	application publication no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent no. 6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246	(2) Obviousness-type double patenting, U.S. patent application no. 11/054,848	D 1
	July 30, 2009	§ 103 (a) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888, U.S. patent application publication no. 2004/0180351, U.S. patent	(1) Obviousness-type double patenting, U.S. patent application no. 10/860,265;	
	December 8, 2008	6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246 § 103 (a) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888, U.S. patent application publication no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent application publication no. 6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246	patent application no. 11/054,848 (1) Obviousness-type double patenting, U.S. patent application no. 10/860,265; (2) Obviousness-type double patenting, U.S. patent application no. 11/054,848	
		no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent no.	10/860,265; (2) Obviousness-type double patenting, U.S.	

	November 30, 2007	patent publication application no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent no. 6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246 § 103 (a) Elbashir, <i>EMBO J.</i> , 2001, 20, 6877-6888, U.S. patent application publication no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent application publication no. 6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246	(2) Obviousness-type double patenting, U.S. patent application no. 10/701,007; (3) Obviousness-type double patenting, U.S. patent application no. 11/054,848 (1) Obviousness-type double patenting, U.S. patent application no. 11/054,848; and (2) Obviousness-type double patenting, U.S. patent application no. 10/701,007	
	June 19, 2008	§ 103 (a) Elbashir, EMBO J., 2001, 20, 6877-6888, U.S. patent application publication no. 2003/0143732, U.S. patent application publication no. 2003/0206887, U.S. patent no. 6,262,036, U.S. patent application publication no. 2005/0142535, and U.S. patent no. 6,133,246	(1) Obviousness-type double patenting, U.S. patent application no. 11/054,848; and (2) Obviousness-type double patenting, U.S. patent application no. 10/701,007	
10/859,825 ISIS-5481	October 3, 2007	§ 102: Damha, <i>J. Am. Chem. Soc.</i> 120: 12976-12977; US 2003/0143732	(1) claim of priority denied; double patenting: 10/700,689; noted that	Pending

			double patenting "may be	
			appropriate" for:	
			10/860,455; 10/909,125;	
			10/664,639; 10/701,007,	
			' ' '	
			10/701,264; 10/701,285,	
			10/936,273; 10/561,618;	
			10/700,689; 10/700,697;	
			10/701,236; 10/701,265;	
			10/701,306; 10/860,265;	
			11/054,848; 10/561,324;	
			11/226,882; 11/565,781;	
			11/565,770; 11/565,773;	
			11/565,794; 11/565,799;	
			11/565,804; 11/565,817;	
			11/565,823; 11/565,833;	
			11/565,839; 11/565,858;	
			11/565,816; 11/565,841;	
			11/569,929; 11/747,042;	
			11/569,931; 11/569,939;	
			11/569,941; 11/569,955	
Jul	ıly 18, 2008	§ 102 (a) U.S. patent	Claim of priority denied	
		application publication no.		
		2003/0143732		
Ma	Tarch 19, 2009	§ 103 (a) Lee, et al., Cell, 1993,	Obviousness-type double	
		75, 843-854; Manche, et al.,	patenting, U.S. patent no.	
		Mol. Cell Biol., 1992, 12,	5,898,031	
		5238-5248; PCT patent		
		application publication no. WO		
		94/01550; and U.S. patent no.		
		5,801,154		